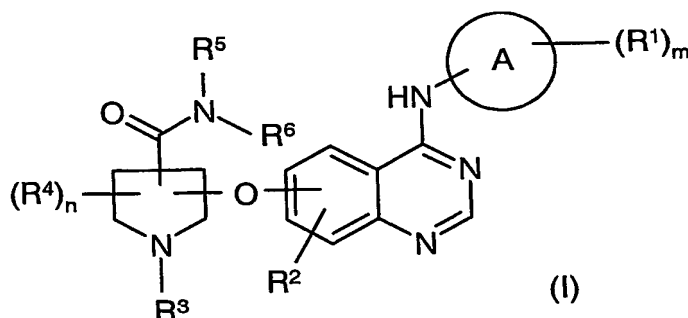


CLAIMS

1. A quinazoline derivative of the Formula (I):



5

wherein:

either R^2 is in the 6-position and the substituted-pyrrolidinyloxy group is in the 7-position of the quinazoline ring or R^2 is in the 7-position and the substituted-pyrrolidinyloxy group is in the 6-position of the quinazoline ring;

A is phenyl or pyridyl;

each R^1 is a substituent on a ring carbon atom in ring A and is independently selected from halogeno, cyano, nitro, hydroxy, carboxy, trifluoromethyl, (1-6C)alkyl, (2-8C)alkenyl, (2-8C)alkynyl, (1-6C)alkoxy, (2-6C)alkenyloxy, (2-6C)alkynyloxy, (1-6C)alkylthio, (1-6C)alkylsulfinyl, (1-6C)alkylsulfonyl, (1-6C)alkoxycarbonyl, ureido, N -(1-6C)alkylureido, N,N -di-[(1-6C)alkyl]ureido, $-NR^aR^b$, $-SO_2NR^aR^b$ and a group of the formula $-CONR^aR^b$ (wherein R^a is hydrogen or (1-6C)alkyl and R^b selected from hydrogen, (1-6C)alkyl, phenyl, benzyl, heterocyclyl, heterocyclyl(1-3C)alkyl, heteroaryl, heteroaryl(1-3C)alkyl, (3-7)cycloalkyl and (3-7)cycloalkyl(1-3C)alkyl wherein any alkyl, heterocyclyl, heteroaryl and cycloalkyl groups in R^a and R^b are optionally substituted by 1, 2 or 3 substituents selected from (1-4C)alkyl, halogeno, hydroxy and (1-4C)alkoxy;

or R^a and R^b together with the nitrogen atom to which they are attached form a 4, 5 or 6-membered ring which optionally contains an additional ring heteroatom selected from nitrogen, oxygen and sulphur and which is optionally substituted by 1 or 2 substituents on an available ring carbon atom, independently selected from halogeno, hydroxy, (1-4C)alkyl and (1-3C)alkylenedioxy and optionally substituted on any available ring nitrogen by a substituent selected from (1-4C)alkyl and (2-4C)alkanoyl (provided the ring is not thereby quaternised),

and wherein any (1-4C)alkyl or (2-4C)alkanoyl group present as a substituent on the ring formed by R^a and R^b together with the nitrogen atom to which they are attached is optionally substituted by 1, 2 or 3 substituents independently selected from halogeno, hydroxyl, (1-4C)alkyl and (1-4C)alkoxy;

- 5 or, when two R¹ groups are attached to adjacent carbon atoms, they may, together with the carbon atoms to which they are attached, form a pyrrole ring, wherein the pyrrole ring is optionally substituted by 1 or 2 substituents independently selected from (1-6C)alkyl, halogeno, cyano, nitro, hydroxy, amino, carbamoyl, sulfamoyl and trifluoromethyl;
or, when two R¹ groups are attached to adjacent carbon atoms, they may, together form a
10 (1-3C)alkylenedioxy group [-O(CH₂)₁₋₃O];
m is 0, 1, 2 or 3;
each R² is selected from hydrogen, (1-6C)alkyl, (3-6C)cycloalkyl, (3-6C)cycloalkyl(1-3C)alkyl and a group of the formula R⁷O-, wherein R⁷ is (1-6C)alkyl optionally substituted by 1, 2 or 3 substituents independently selected from hydroxy and a group of the formula R⁸O- (wherein R⁸

15 is (1-3C)alkyl);

R³ is selected from hydrogen, (1-6C)alkyl, (3-6C)cycloalkyl, (3-6C)cycloalkyl(1-3C)alkyl, (1-6C)alkylthio, (1-6C)alkylsulfinyl, (1-6C)alkylsulfonyl, (2-6C)alkanoyl, carbamoyl(1-6C)alkyl, N-(1-6C)alkylcarbamoyl(1-6C)alkyl, N,N-di-[(1-6C)alkyl]carbamoyl(1-6C)alkyl, sulfamoyl(1-6C)alkyl,

- 20 N-(1-6C)alkylsulfamoyl(1-6C)alkyl, N,N-di-[(1-6C)alkyl]sulfamoyl(1-6C)alkyl and (2-6C)alkanoyl(1-6C)alkyl,

and wherein any (1-6C)alkyl or (2-6C)alkanoyl group within R³ is optionally substituted by 1, 2 or 3 substituents independently selected from halogeno, hydroxy and (1-6C)alkyl and/or optionally a substituent selected from cyano, nitro, (2-8C)alkenyl,

- 25 (2-8C)alkynyl, (1-6C)alkoxy and NR^cR^d, wherein R^c is hydrogen or (1-4C)alkyl and R^d is hydrogen or (1-4C)alkyl, and wherein any (1-4C)alkyl in R^c or R^d is optionally substituted by 1, 2 or 3 substituents independently selected from halogeno and hydroxy and/or optionally a substituent selected from cyano, nitro and (1-4C)alkoxy,

- or R^c and R^d together with the nitrogen atom to which they are attached form a 4, 5 or
30 6 membered ring which optionally contains an additional ring heteroatom selected from nitrogen, oxygen and sulphur and which is optionally substituted by 1 or 2 substituents on an available ring carbon atom, independently selected from halogeno, hydroxy, (1-4C)alkyl and

(1-3C)alkylenedioxy, and optionally substituted on any available ring nitrogen by a substituent selected from (1-4C)alkyl and (2-4C)alkanoyl (provided the ring is not thereby quaternised),

and wherein any (1-4C)alkyl or (2-4C)alkanoyl group present as a substituent on the ring formed by R^c and R^d together with the nitrogen atom to which they are attached is

5 optionally substituted by 1, 2 or 3 substituents independently selected from halogeno and hydroxy and/or optionally a substituent selected from (1-4C)alkyl and (1-4C)alkoxy;

each R⁴ is independently selected from (1-4C)alkyl, (1-4C)alkoxy, cyano, halogeno, hydroxyl and oxo;

n is 0, 1 or 2;

10 R⁵ is hydrogen or (1-6C)alkyl;

R⁶ is selected from hydrogen, (1-6C)alkyl, (2-6C)alkenyl, (2-6C)alkynyl, (1-6C)alkoxy, (3-7)cycloalkyl, (1-6C)alkylsulfonyl, heterocyclyl, heteroaryl, (3-7)cycloalkyl(1-3C)alkyl, (3-7)heterocyclyl(1-3C)alkyl and heteroaryl(1-3C)alkyl,

and wherein any (1-3C)alkyl, (1-6C)alkyl, (3-7)cycloalkyl, heteroaryl or heterocyclyl group

15 within R⁵ or R⁶ is optionally substituted (on any available carbon atoms) by 1, 2 or 3 substituents independently selected from halogeno, hydroxy(1-6C)alkyl, (1-6C)alkoxycarbonyl, carbamoyl, (2-6C)alkanoylamino and hydroxy and/or optionally a substituent selected from oxo, cyano, nitro and (1-4C)alkoxy,

and wherein any heterocyclyl group within R⁶ is optionally substituted on any available ring

20 nitrogen (provided the ring is not thereby quaternised) by (1-4C)alkyl or (2-4C)alkanoyl, or R⁵ and R⁶ together with the nitrogen atom to which they are attached form a 4, 5 or 6

membered ring which is optionally substituted by 1 or 2 substituents on an available ring carbon atom, independently selected from halogeno, hydroxy, (1-4C)alkyl and

(1-3C)alkylenedioxy, and optionally substituted on any available ring nitrogen by a substituent

25 selected from (1-4C)alkyl and (2-4C)alkanoyl (provided the ring is not thereby quaternised),

and wherein any (1-4C)alkyl or (2-4C)alkanoyl group present as a substituent on the

ring formed by R⁵ and R⁶ together with the nitrogen atom to which they are attached is

optionally substituted by 1, 2 or 3 substituents independently selected from halogeno and hydroxy and/or optionally a substituent selected from (1-4C)alkyl and (1-4C)alkoxy;

30 provided that when the pyrrolidinyloxy group is linked to the 6-position of the

quinazoline ring, m is 2 and substituents R¹ are both halogeno and attached to the 2- and 3-

positions of the ring A, then R⁶ is selected from substituted-(1-6C)alkyl (wherein substituted-

- (1-6C)alkyl is (1-6C)alkyl substituted by 1, 2 or 3 substituents independently selected from halogeno, hydroxy(1-6C)alkyl, (1-6C)alkoxycarbonyl, carbamoyl, (2-6C)alkanoylamino, (1-6C)alkylamino, di-[(1-6C)alkyl]amino and hydroxy and/or optionally a substituent selected from oxo, cyano, nitro and (1-4C)alkoxy), (2-6C)alkenyl, (2-6C)alkynyl, (1-6C)alkoxy, (3-7)cycloalkyl, (1-6C)alkylsulfonyl, (3-7)heterocyclyl, heteroaryl, (3-7)cycloalkyl(1-6C)alkyl, (3-7)heterocyclyl(1-6C)alkyl and heteroaryl(1-6C)alkyl, and wherein any (3-7)cycloalkyl, heteroaryl or (3-7)heterocyclyl group within R⁶ is optionally substituted (on any available carbon atoms) by 1, 2 or 3 substituents independently selected from halogeno, hydroxy, (1-6C)alkyl, hydroxy(1-6C)alkyl, (1-6C)alkoxycarbonyl, carbamoyl, (2-6C)alkanoylamino and hydroxy and/or optionally a substituent selected from oxo, cyano, nitro and (1-4C)alkoxy, and wherein any heteroaryl or heterocyclyl group within R⁶ is optionally substituted on any available ring nitrogen (provided the ring is not thereby quaternised) by (1-4C)alkyl or (2-4C)alkanoyl, or
- 15 R⁵ and R⁶ together with the nitrogen atom to which they are attached form a 4, 5 or 6 membered ring which contains one or two nitrogen atoms as the only heteroatoms present in the ring and which is optionally substituted on an available ring carbon atom by 1 or 2 substituents independently selected from hydroxy, carbamoyl, (1-4C)alkyl, and (1-3C)alkylenedioxy; and wherein any 4, 5 or 6 membered heterocyclic ring formed by R⁵ and R⁶ is optionally substituted on any available ring nitrogen (provided the ring is not thereby quaternised) by (1-4C)alkyl or (2-4C)alkanoyl;
- 20 or a pharmaceutically-acceptable salt thereof.

2. A quinazoline derivative according to claim 1, wherein R⁵ is hydrogen or (1-6C)alkyl and R⁶ is selected from hydrogen, (1-6C)alkyl, (2-6C)alkenyl, (2-6C)alkynyl, (1-6C)alkoxy, (3-7)cycloalkyl, (1-6C)alkylsulfonyl, heterocyclyl, heteroaryl, (3-7)cycloalkyl(1-3C)alkyl, (3-7)heterocyclyl(1-3C)alkyl and heteroaryl(1-3C)alkyl, and wherein any (1-3C)alkyl, (1-6C)alkyl, (3-7)cycloalkyl, heteroaryl or heterocyclyl group within R⁵ or R⁶ is optionally substituted (on any available carbon atoms) by 1, 2 or 3 substituents independently selected from halogeno, hydroxy(1-6C)alkyl, (1-6C)alkoxycarbonyl, carbamoyl, (2-6C)alkanoylamino and hydroxy and/or optionally a substituent selected from oxo, cyano, nitro and (1-4C)alkoxy,
- 30

and wherein any heterocyclyl group within R⁶ is optionally substituted on any available ring nitrogen (provided the ring is not thereby quaternised) by (1-4C)alkyl or (2-4C)alkanoyl, or R⁵ and R⁶ together with the nitrogen atom to which they are attached form a 4, 5 or 6 membered ring which is optionally substituted by 1 or 2 substituents on an available ring carbon atom, independently selected from halogeno, hydroxy, (1-4C)alkyl and (1-3C)alkylenedioxy, and optionally substituted on any available ring nitrogen by a substituent selected from (1-4C)alkyl and (2-4C)alkanoyl (provided the ring is not thereby quaternised), and wherein any (1-4C)alkyl or (2-4C)alkanoyl group present as a substituent on the ring formed by R⁵ and R⁶ together with the nitrogen atom to which they are attached is optionally substituted by 1, 2 or 3 substituents independently selected from halogeno and hydroxy and/or optionally a substituent selected from (1-4C)alkyl and (1-4C)alkoxy; provided that when the pyrrolidinyloxy group is linked to the 6-position of the quinazoline ring, m is 2 and substituents R¹ are both halogeno and attached to the 2- and 3-positions of the ring A, then R⁶ is selected from substituted-(1-6C)alkyl (wherein substituted-(1-6C)alkyl is (1-6C)alkyl substituted by 1, 2 or 3 substituents independently selected from (1-6C)alkoxycarbonyl, carbamoyl, (2-6C)alkanoylamino, and oxo or a (1-6C)alkoxycarbonyl together with a hydroxy group), (1-6C)alkoxy, (1-6C)alkylsulfonyl, (3-7)heterocyclyl (wherein the heterocyclyl is carbon linked), heteroaryl, (3-7)heterocyclyl(1-6C)alkyl (wherein the heterocyclyl is carbon linked to the (1-6C)alkyl moiety) and heteroaryl(1-6C)alkyl, and wherein any heteroaryl or (3-7)heterocyclyl group within R⁶ is optionally substituted (on any available carbon atoms) by 1, 2 or 3 substituents independently selected from halogeno, (1-6C)alkyl, hydroxy(1-6C)alkyl, (1-6C)alkoxycarbonyl, carbamoyl, (2-6C)alkanoylamino and hydroxy and/or optionally a substituent selected from oxo, cyano, nitro and (1-4C)alkoxy, and wherein any heteroaryl or heterocyclyl group within R⁶ is optionally substituted on any available ring nitrogen (provided the ring is not thereby quaternised) by (1-4C)alkyl or (2-4C)alkanoyl, or R⁵ and R⁶ together with the nitrogen atom to which they are attached form a 4, 5 or 6 membered ring which contains one or two nitrogen atoms as the only heteroatoms present in the ring and which is substituted on an available ring carbon atom by 1 or 2 substituents independently selected from carbamoyl and (1-3C)alkylenedioxy.

3. A quinazoline derivative according to claim 1 or claim 2, wherein R⁵ is hydrogen, methyl, ethyl propyl, isopropyl or isobutyl and R⁶ is selected from hydrogen, methyl, ethyl propyl, isopropyl, isobutyl, vinyl, isopropenyl, allyl, but-2-enyl ethynyl, 2-propynyl, butynyl, methoxy, ethoxy propoxy, isopropoxy, cyclopropyl, cyclopentyl, cyclohexyl, azetidinyl, oxazepanyl, pyrrolinyl, pyrrolidinyl, morpholinyl, tetrahydro-1,4-thiazinyl, piperidinyl, homopiperidinyl, piperazinyl, homopiperazinyl, dihydropyridinyl, tetrahydropyridinyl, dihydropyrimidinyl, tetrahydropyrimidinyl, tetrahydrothienyl, tetrahydrothiopyranyl, thiomorpholinyl, pyrazolyl, thienyl, oxazolyl, isoxazolyl, imidazolyl, pyridinyl, pyridazinyl, pyrazinyl, pyrimidyl, furanyl, pyrazolyl, thiazolyl, isothiazolyl, thiadiazolyl, cyclopropylmethyl, cyclopentylmethyl, cyclohexylmethyl, 2-cyclopropylethyl, 2-cyclopentylethyl, 2-cyclohexylethyl, azetidinylmethyl, oxazepanylmethyl, pyrrolinylmethyl, pyrrolidinylmethyl, morpholinylmethyl, tetrahydro-1,4-thiazinylmethyl, piperidinylmethyl, homopiperidinylmethyl, piperazinylmethyl, homopiperazinylmethyl, dihydropyridinylmethyl, tetrahydropyridinylmethyl, dihydropyrimidinylmethyl, tetrahydropyrimidinylmethyl, tetrahydrothienylmethyl, tetrahydrothiopyranyl, thiomorpholinylmethyl, pyrazolylmethyl, thienylmethyl, oxazolylmethyl, isoxazolylmethyl, imidazolylmethyl, pyridinylmethyl, pyridazinylmethyl, pyrazinylmethyl, pyrimidylmethyl, furanyl, pyrazolylmethyl, thiazolylmethyl, isothiazolylmethyl, thiadiazolylmethyl, 2-(azetidiny)ethyl, 2-(oxazepanyl)ethyl, 2-(pyrrolinyl)ethyl, 2-(pyrrolidinyl)ethyl, 2-(morpholinyl)ethyl, 2-(tetrahydro-1,4-thiazinyl)ethyl, 2-(piperidinyl)ethyl, 2-(homopiperidinyl)ethyl, 2-(piperazinyl)ethyl, 2-(homopiperazinyl)ethyl, 2-(dihydropyridinyl)ethyl, 2-(tetrahydropyridinyl)ethyl, 2-(dihydropyrimidinyl)ethyl, 2-(tetrahydropyrimidinyl)ethyl, 2-(tetrahydrothienyl)ethyl, 2-(tetrahydrothiopyranyl)ethyl, 2-(thiomorpholinyl)ethyl, 2-(pyrazolyl)ethyl, 2-(thienyl)ethyl, 2-(oxazolyl)ethyl, 2-(isoxazolyl)ethyl, 2-(imidazolyl)ethyl, 2-(pyridinyl)ethyl, 2-(pyridazinyl)ethyl, 2-(pyrazinyl)ethyl, 2-(pyrimidyl)ethyl, 2-(furanyl)ethyl, 2-(pyrazolyl)ethyl, 2-(thiazolyl)ethyl, 2-(isothiazolyl)ethyl and 2-(thiadiazolyl)ethyl, and wherein any alkyl, cycloalkyl, heteroaryl or heterocyclyl group within R⁵ or R⁶ is optionally substituted (on any available carbon atoms) by 1 or 2 substituents independently selected from fluoro, chloro, bromo, hydroxymethyl, 2-hydroxyethyl, methoxycarbonyl, ethoxycarbonyl, carbamoyl, acetamido, propionamido and hydroxy and/or optionally a substituent selected from oxo, cyano, methoxy and ethoxy,

and wherein any heterocyclyl group within R⁶ is optionally substituted on any available ring nitrogen (provided the ring is not thereby quaternised) by methyl, ethyl, acetyl or propionyl, or R⁵ and R⁶ together with the nitrogen atom to which they are attached form a azetidin-1-yl, pyrrolin-1-yl, pyrrolidin-1-yl, piperidino, morpholino or piperazino ring which is optionally substituted by 1 or 2 substituents on an available ring carbon atom, independently selected from fluoro, chloro, bromo, hydroxy, methyl, ethyl and propylenedioxy, and optionally substituted on any available ring nitrogen by a substituent selected from methyl, ethyl, acetyl and propionyl (provided the ring is not thereby quaternised),

and wherein any alkyl or alkanoyl group present as a substituent on the ring formed by R⁵ and R⁶ together with the nitrogen atom to which they are attached is optionally substituted by 1 or 2 substituents independently selected from fluoro, chloro, bromo and hydroxy and/or optionally a substituent selected from methyl, ethyl, methoxy and ethoxy; provided that when the pyrrolidinyloxy group is linked to the 6-position of the quinazoline ring, m is 2 and substituents R¹ are both halogeno and attached to the 2- and 3- positions of the ring A, then R⁶ is selected from substituted-methyl, substituted-ethyl substituted-propyl, substituted-isopropyl, substituted-isobutyl, (wherein the substituted groups are substituted by 1 or 2 substituents independently selected from methoxycarbonyl, ethoxycarbonyl, carbamoyl, acetamido, propionamido and oxo or a methoxycarbonyl group together with a hydroxy group or an ethoxycarbonyl group together with a hydroxy group) methoxy, ethoxy, propoxy, isopropoxy,

a carbon linked heterocyclyl group selected from azetidiny, oxazepanyl, pyrrolinyl, pyrrolidinyl, morpholinyl, tetrahydrofuranyl, tetrahydro-1,4-thiazinyl, piperidinyl, homopiperidinyl, piperazinyl, homopiperazinyl, dihydropyridinyl, tetrahydropyridinyl, dihydropyrimidinyl, tetrahydropyrimidinyl, tetrahydrothienyl, tetrahydropyranyl, tetrahydrothiopyranyl, thiomorpholinyl,

a heteroaryl group selected from pyrazolyl, thienyl, oxazolyl, isoxazolyl, imidazolyl, pyridinyl, pyridazinyl, pyrazinyl, pyrimidyl, furanyl, thiazolyl, isothiazolyl, thiadiazolyl,

a (3-7)heterocyclyl(1-6C)alkyl group (wherein the heterocyclyl is carbon linked to the (1-6C)alkyl moiety) selected from azetidinylmethyl, oxazepanylmethyl, pyrrolinylmethyl, pyrrolidinylmethyl, morpholinylmethyl, tetrahydro-1,4-thiazinylmethyl, piperidinylmethyl, homopiperidinylmethyl, piperazinylmethyl, homopiperazinylmethyl, dihydropyridinylmethyl, tetrahydropyridinylmethyl, dihydropyrimidinylmethyl, tetrahydropyrimidinylmethyl,

- tetrahydrofuranylmethyl, tetrahydrothienylmethyl, tetrahydropyranylmethyl, tetrahydrothiopyranylmethyl, thiomorpholinylmethyl, 2-(azetidiny)ethyl, 2-(oxazepanyl)ethyl, 2-(pyrrolinyl)ethyl, 2-(pyrrolidinyl)ethyl, 2-(morpholinyl)ethyl, 2-(tetrahydro-1,4-thiazinyl)ethyl, 2-(piperidinyl)ethyl, 2-(homopiperidinyl)ethyl, 2-(piperazinyl)ethyl, 2-(homopiperazinyl)ethyl, 2-(dihydropyridinyl)ethyl, 2-(tetrahydropyridinyl)ethyl, 2-(dihydropyrimidinyl)ethyl, 2-(tetrahydropyrimidinyl)ethyl, 2-(tetrahydrofuranylmethyl), 2-(tetrahydrothienyl)ethyl, 2-(tetrahydropyranylmethyl), 2-(tetrahydrothiopyranylmethyl), 2-(thiomorpholinyl)ethyl, a heteroaryl(1-6C)alkyl group selected from pyrazolylmethyl, thienylmethyl, oxazolylmethyl, isoxazolylmethyl, imidazolylmethyl, pyridinylmethyl, pyridazinylmethyl, pyrazinylmethyl, pyrimidinylmethyl, furanylmethyl, pyrazolylmethyl, thiazolylmethyl, isothiazolylmethyl, thiadiazolylmethyl, 2-(pyrazolyl)ethyl, 2-(thienyl)ethyl, 2-(oxazolyl)ethyl, 2-(isoxazolyl)ethyl, 2-(imidazolyl)ethyl, 2-(pyridinyl)ethyl, 2-(pyridazinyl)ethyl, 2-(pyrazinyl)ethyl, 2-(pyrimidinyl)ethyl, 2-(furanylmethyl), 2-(pyrazolyl)ethyl, 2-(thiazolyl)ethyl, 2-(isothiazolyl)ethyl and 2-(thiadiazolyl)ethyl, and wherein any heteroaryl or heterocyclyl group within R⁶ is optionally substituted (on any available carbon atoms) by 1 or 2 substituents independently selected from fluoro, chloro, bromo, hydroxymethyl, 2-hydroxyethyl, methoxycarbonyl, ethoxycarbonyl, carbamoyl, acetamido, propionamido and hydroxy and/or optionally a substituent selected from oxo, cyano, methoxy and ethoxy, and wherein any heteroaryl or heterocyclyl group within R⁶ is optionally substituted on any available ring nitrogen (provided the ring is not thereby quaternised) by methyl, ethyl, acetyl or propionyl or R⁵ and R⁶ together with the nitrogen atom to which they are attached form an azetidin-1-yl ring substituted carbamoyl or (1-3C)alkylenedioxy.

4. A quinazoline derivative according to any one of claims 1 to 3, wherein R⁵ is hydrogen, methyl or ethyl and R⁶ is selected from hydrogen, methyl, ethyl, propyl, isopropyl, isobutyl, vinyl, isoprop-2-enyl, allyl, but-2-enyl ethynyl, 2-prop-2-ynyl, but-3-ynyl, methoxy, ethoxy, cyclopropyl, cyclopentyl, cyclohexyl, azetidiny, pyrrolinyl, pyrrolidinyl, morpholinyl, piperidinyl, piperazinyl, tetrahydropyridinyl, thiomorpholinyl, 1,2,3,6-tetrahydropyridin-1-yl, pyrazolyl, thienyl, oxazolyl, isoxazolyl, imidazolyl, pyridinyl, pyridazinyl, pyrazinyl, pyrimidinyl,

furanyl, pyrazolyl, thiazolyl, isothiazolyl, cyclopropylmethyl, cyclopentylmethyl, cyclohexylmethyl, 2-cyclopropylethyl, 2-cyclopentylethyl, 2-cyclohexylethyl, azetidylmethyl, pyrrolinylmethyl, pyrrolidinylmethyl, morpholinylmethyl, piperidinylmethyl, piperazinylmethyl, tetrahydropyridinylmethyl, thiomorpholinylmethyl, pyrazolylmethyl, thienylmethyl,

- 5 oxazolylmethyl, isoxazolylmethyl, imidazolylmethyl, pyridinylmethyl, pyridazinylmethyl, pyrazinylmethyl, pyrimidinylmethyl, furanylmethyl, pyrazolylmethyl, thiazolylmethyl, isothiazolylmethyl, 2-(azetidyl)ethyl, 2-(pyrrolinyl)ethyl, 2-(pyrrolidinyl)ethyl, 2-(morpholinyl)ethyl, 2-(piperidinyl)ethyl, 2-(piperazinyl)ethyl, 2-(tetrahydropyridinyl)ethyl, 2-(thiomorpholinyl)ethyl, 2-(pyrazolyl)ethyl, 2-(thienyl)ethyl, 2-(oxazolyl)ethyl, 2-(isoxazolyl)ethyl, 2-(imidazolyl)ethyl, 2-(pyridinyl)ethyl, 2-(pyridazinyl)ethyl, 2-(pyrazinyl)ethyl, 2-(pyrimidinyl)ethyl, 2-(furanyl)ethyl, 2-(pyrazolyl)ethyl, 2-(thiazolyl)ethyl and 2-(isothiazolyl)ethyl,

and wherein any alkyl, cycloalkyl, heteroaryl or heterocyclyl group within R⁵ or R⁶ is optionally substituted (on any available carbon atoms) by 1 or 2 substituents independently

- 15 selected from fluoro, chloro, bromo, hydroxymethyl, 2-hydroxyethyl, methoxycarbonyl, ethoxycarbonyl, carbamoyl, acetamido and hydroxy and/or optionally a substituent selected from oxo, cyano, methoxy and ethoxy,

and wherein any heterocyclyl group within R⁶ is optionally substituted on any available ring nitrogen (provided the ring is not thereby quaternised) by methyl, ethyl, acetyl or propionyl, or

- 20 R⁵ and R⁶ together with the nitrogen atom to which they are attached form a azetidin-1-yl, pyrrolin-1-yl, pyrrolidin-1-yl, piperidino, morpholino or piperazino ring which is optionally substituted by 1 or 2 substituents on an available ring carbon atom, independently selected from fluoro, chloro, hydroxy, methyl, ethyl and propylenedioxy, and optionally substituted on any available ring nitrogen by a substituent selected from methyl, ethyl, acetyl and propionyl
25 (provided the ring is not thereby quaternised),

and wherein any alkyl or alkanoyl group present as a substituent on the ring formed by R⁵ and R⁶ together with the nitrogen atom to which they are attached is optionally substituted by 1 or 2 substituents independently selected from fluoro, chloro and hydroxy and/or optionally a substituent selected from methyl, ethyl, methoxy and ethoxy;

- 30 provided that when the pyrrolidinyl group is linked to the 6-position of the quinazoline ring, m is 2 and substituents R¹ are both halogeno and attached to the 2- and 3- positions of the ring A, then R⁶ is selected from substituted-methyl, substituted-ethyl substituted-propyl,

substituted-isopropyl, substituted-isobutyl, (wherein the substituted groups are substituted by 1 or 2 substituents independently selected from methoxycarbonyl, ethoxycarbonyl, carbamoyl, acetamido and oxo or a methoxycarbonyl group together with a hydroxy group), methoxy, ethoxy,

5 a carbon linked heterocyclyl group selected from azetidiny, pyrroliny, pyrrolidinyl, morpholinyl, tetrahydrofuranyl, piperidiny, piperazinyl, tetrahydropyridiny, tetrahydropyranyl, thiomorpholinyl,

a heteroaryl group selected from pyrazolyl, thienyl, oxazolyl, isoxazolyl, imidazolyl, pyridiny, pyridazinyl, pyrazinyl, pyrimidyl, furanyl, pyrazolyl, thiazolyl, isothiazolyl,

10 a (3-7)heterocyclyl(1-6C)alkyl group (wherein the heterocyclyl is carbon linked to the (1-6C)alkyl moiety) selected from azetidinylmethyl, pyrrolinylmethyl, pyrrolidinylmethyl, morpholinylmethyl, piperidinylmethyl, piperazinylmethyl, tetrahydrofuranylmethyl, tetrahydropyranylmethyl, tetrahydropyridinylmethyl, thiomorpholinylmethyl, 2-(azetidiny)ethyl, 2-(pyrroliny)ethyl, 2-(pyrrolidinyl)ethyl, 2-(morpholinyl)ethyl, 2-(piperidiny)ethyl, 2-(piperazinyl)ethyl, 2-(tetrahydrofuranyl)ethyl, 2-(tetrahydropyranyl)methyl, 2-(tetrahydropyridiny)ethyl, 2-(thiomorpholinyl)ethyl,

15 a heteroaryl(1-6C)alkyl group selected from pyrazolylmethyl, thienylmethyl, oxazolylmethyl, isoxazolylmethyl, imidazolylmethyl, pyridinylmethyl, pyridazinylmethyl, pyrazinylmethyl, pyrimidylmethyl, furanylmethyl, pyrazolylmethyl, thiazolylmethyl, isothiazolylmethyl, 2-(pyrazolyl)ethyl, 2-(thienyl)ethyl, 2-(oxazolyl)ethyl, 2-(isoxazolyl)ethyl, 2-(imidazolyl)ethyl, 2-(pyridiny)ethyl, 2-(pyridazinyl)ethyl, 2-(pyrazinyl)ethyl, 2-(pyrimidyl)ethyl, 2-(furanyl)ethyl, 2-(pyrazolyl)ethyl, 2-(thiazolyl)ethyl and 2-(isothiazolyl)ethyl,

25 and wherein any heteroaryl or heterocyclyl group within R⁶ is optionally substituted (on any available carbon atoms) by 1 or 2 substituents independently selected from fluoro, chloro, bromo, hydroxymethyl, 2-hydroxyethyl, methoxycarbonyl, ethoxycarbonyl, carbamoyl, acetamido and hydroxy and/or optionally a substituent selected from oxo, cyano, methoxy and ethoxy,

30 and wherein any heteroaryl or heterocyclyl group within R⁶ is optionally substituted on any available ring nitrogen (provided the ring is not thereby quaternised) by methyl, ethyl, acetyl or propionyl;

or R⁵ and R⁶ together with the nitrogen atom to which they are attached form an azetidin-1-yl ring substituted by a carbamoyl group.

5. A quinazoline derivative according to any one of claims 1 to 4, wherein R⁵ is hydrogen or methyl and R⁶ is selected from hydrogen, methyl, ethyl, propyl, isopropyl, vinyl, isoprop-2-enyl, allyl, but-2-enyl ethynyl, 2-propynyl, but-3-ynyl, methoxy, cyclopropyl, cyclopentyl, 1-(hydroxymethyl)cyclopentyl, cyclohexyl, 4-hydroxycyclohexyl, cyclopropylmethyl, cyclopentylmethyl, methoxymethyl, 2-(methoxy)ethyl, 2-(ethoxy)ethyl, carbamoylmethyl, 2-(acetyl)ethyl, cyanomethyl, 2-(cyano)ethyl, 2,3-dihydroxypropyl, 2-(hydroxyl)-1,1-dimethylethyl, 2,2,2-trifluoroethyl, 1-(ethoxycarbonyl)-2-hydroxyethyl, 2-acetamido)ethyl, tetrahydrofuran-2-ylmethyl, imidazol-2-ylmethyl, 1-methylpyrazol-5-yl, 1-methylpyrazol-5-yl, 3-methylpyrazol-5-yl, imidazol-1-ylmethyl, 2-(imidazol-1-yl)ethyl, furan-2-ylmethyl, 2-(furan-2-yl)ethyl, 5-methylisoxazol-3-ylmethyl, thien-3yl, morpholino, piperidin-4-yl, 1-methylpiperidin-4-yl, tetrahydro-2H-pyran-4-yl and 3-oxotetrahydrofuran-4-yl,
- 15 or R⁵ and R⁶ together with the nitrogen atom to which they are attached form a 3-hydroxyazetidin-1-yl, 2-carbamoylazetidin-1-yl, pyrrolin-1-yl, pyrrolidin-1-yl, 3-hydroxy, pyrrolidin-1-yl, piperidino, morpholino or piperazino group; provided that when the pyrrolidinyloxy group is linked to the 6-position of the quinazoline ring, m is 2 and substituents R¹ are both halogeno and attached to the 2- and 3- positions of the ring A, then R⁶ is selected from methoxy, carbamoylmethyl, 2-(hydroxy)-1-(methoxycarbonyl)ethyl, 1-(ethoxycarbonyl)-2-hydroxyethyl, 2-(acetamido)ethyl, piperidin-4-yl, 1-methylpiperidin-4-yl, tetrahydropyran-4-yl, 4-hydroxytetrahydrofuran-3-yl, 3-oxotetrahydrofuran-4-yl, 1-methylpyrazol-5-yl, thien-3yl, 3-methylpyrazol-5-yl, tetrahydrofuran-2-ylmethyl, tetrahydropyran-4-ylmethyl, furan-2-ylmethyl, 2-(furan-2-yl)ethyl, imidazol-1-ylmethyl, imidazol-2-ylmethyl, imidazol-2-ylmethyl, 2-(imidazol-1-yl)ethyl, 2-(imidazol-4-yl)ethyl and 5-methylisoxazol-3-ylmethyl or R⁵ and R⁶ together with the nitrogen atom to which they are attached form an azetidinyl substituted in the 2 position by a carbamoyl group.

6. A quinazoline derivative according to claim 1 or claim 2, wherein R⁵ is hydrogen or (1-6C)alkyl and R⁶ is selected from hydrogen, (1-6C)alkyl, (2-6C)alkenyl, (2-6C)alkynyl, (1-6C)alkoxy, (3-7)cycloalkyl, (1-6C)alkylsulfonyl, heterocyclyl, heteroaryl, (3-7)cycloalkyl(1-3C)alkyl, (3-7)heterocyclyl(1-3C)alkyl and heteroaryl(1-3C)alkyl,

and wherein any (1-3C)alkyl, (1-6C)alkyl, (3-7)cycloalkyl, heteroaryl or heterocyclyl group within R⁵ or R⁶ is optionally substituted (on any available carbon atoms) by 1, 2 or 3 substituents independently selected from halogeno, hydroxy(1-6C)alkyl, (1-6C)alkoxycarbonyl, carbamoyl, (2-6C)alkanoylamino and hydroxy and/or optionally a substituent selected from

5 oxo, cyano, nitro and (1-4C)alkoxy,

and wherein any heterocyclyl group within R⁶ is optionally substituted on any available ring nitrogen (provided the ring is not thereby quaternised) by (1-4C)alkyl or (2-4C)alkanoyl, or R⁵ and R⁶ together with the nitrogen atom to which they are attached form a 4, 5 or 6

10 membered ring which is optionally substituted by 1 or 2 substituents on an available ring carbon atom, independently selected from halogeno, hydroxy, (1-4C)alkyl and (1-3C)alkylenedioxy, and optionally substituted on any available ring nitrogen by a substituent selected from (1-4C)alkyl and (2-4C)alkanoyl (provided the ring is not thereby quaternised),

and wherein any (1-4C)alkyl or (2-4C)alkanoyl group present as a substituent on the ring formed by R⁵ and R⁶ together with the nitrogen atom to which they are attached is

15 optionally substituted by 1, 2 or 3 substituents independently selected from halogeno and hydroxy and/or optionally a substituent selected from (1-4C)alkyl and (1-4C)alkoxy;

provided that when the pyrrolidinyloxy group is linked to the 6-position of the quinazoline ring, m is 2 and substituents R¹ are both halogeno and attached to the 2- and 3-positions of the ring A, then R⁶ is selected from (3-7)heterocyclyl (wherein the heterocyclyl is

20 carbon linked), heteroaryl, (3-7)heterocyclyl(1-6C)alkyl (wherein the heterocyclyl is carbon linked to the (1-6C)alkyl moiety) and heteroaryl(1-6C)alkyl,

and wherein any heteroaryl or (3-7)heterocyclyl group within R⁶ is optionally substituted (on any available carbon atoms) by 1, 2 or 3 substituents independently selected from halogeno, (1-6C)alkyl, hydroxy(1-6C)alkyl, (1-6C)alkoxycarbonyl, carbamoyl, (2-6C)alkanoylamino and

25 hydroxy and/or optionally a substituent selected from oxo, cyano, nitro and (1-4C)alkoxy,

and wherein any heteroaryl or heterocyclyl group within R⁶ is optionally substituted on any available ring nitrogen (provided the ring is not thereby quaternised) by (1-4C)alkyl or (2-4C)alkanoyl.

30 7. A quinazoline derivative according to anyone of the preceding claims, wherein m is 0, 1, 2 or 3 and R¹ is independently selected from halogeno, cyano, nitro, hydroxy, trifluoromethyl, (1-6C)alkyl, (1-6C)alkoxy, (1-6C)alkylthio, (1-6C)alkylsulfinyl, (1-6C)alkylsulfonyl, ureido, N-(1-6C)alkylureido, N,N-di-[(1-6C)alkyl]ureido, -NR^aR^b,

-SO₂NR^aR^b and a group of the formula -CONR^aR^b (wherein R^a and R^b are as hereinabove defined);

or, when two R¹ groups are attached to adjacent carbon atoms, they may, together with the carbon atoms to which they are attached, form a pyrrole ring, wherein the pyrrole ring is

5 optionally substituted by 1 or 2 substituents independently selected from (1-6C)alkyl, halogeno, cyano, nitro, hydroxy, amino, carbamoyl, sulfamoyl and trifluoromethyl;

or, when two R¹ groups are attached to adjacent carbon atoms, they may, together form a (1-3C)alkylenedioxy group.

10 8. A quinazoline derivative according to claim 7, wherein m is 0, 1 or 2 and R¹ is independently selected from fluoro, chloro, cyano, trifluoromethyl, methyl, methoxy, methylthio, isobutylthio, sulfamoyl, and a group of the formula -CONR^aR^b (wherein R^a is hydrogen or methyl and R^b selected from hydrogen, methyl, ethyl, isobutyl, furanyl, cyclopentyl and cyclohexyl, wherein any alkyl, (3-7)cycloalkyl, heteroaryl in R^a and R^b are

15 optionally substituted by 1 or 2 substituents selected from hydroxy and methoxy;

or R^a and R^b together with the nitrogen atom to which they are attached form a 1,2,3,6-tetrahydropyridin-1-yl, pyrrolidin-1-yl, piperidino, piperazin-1-yl or morpholino ring, which is optionally substituted by 1 or 2 substituents on an available ring carbon atom, independently selected from hydroxyl and optionally substituted on any available ring nitrogen by a

20 substituent selected from methyl and acetyl (provided the ring is not thereby quaternised), or, when two R¹ groups are attached to adjacent carbon atoms, they may, together with the carbon atoms to which they are attached, form a pyrrole ring, wherein the pyrrole ring is optionally substituted by 1 or 2 substituents independently selected from hydroxy; or, when two R¹ groups are attached to adjacent carbon atoms, they may, together form a

25 (1-3C)alkylenedioxy group.

9. A quinazoline derivative according to claim 7 or claim 8, wherein m is 2 and R¹ is positioned in the 2- and 3-positions of ring A and R¹ is independently selected from fluoro and chloro.

30

10. A quinazoline derivative according to any one of the preceding claims, wherein ring A is phenyl or pyrid-3-yl.

11. A quinazoline derivative according to any one of the preceding claims, wherein ring A is phenyl.
- 5 12. A quinazoline derivative according to any one of the preceding claims, wherein R² is selected from hydrogen, (1-6C)alkyl and a group of the formula R⁷O-, wherein R⁷ is (1-6C)alkyl optionally substituted by 1 or 2 substituents independently selected from hydroxy and a group of the formula R⁸O- (wherein R⁸ is (1-3C)alkyl).
- 10 13. A quinazoline derivative according to any one of the preceding claims, wherein R² is selected from hydrogen, methyl, ethyl and a group of the formula R⁷O-, wherein R⁷ is methyl or ethyl.
14. A quinazoline derivative according to any one of the preceding claims, wherein R² is
15 methoxy.
15. A quinazoline derivative according to any one of claims 1 to 13, wherein R² is hydrogen.
- 20 16. A quinazoline derivative according to any one of the preceding claims, wherein R² is in the 6-position and the substituted-pyrrolidinyloxy group is in the 7-position of the quinazoline ring.
17. A quinazoline derivative according to any one of claims 1 to 15, wherein R² is in the 7-
25 position and the substituted-pyrrolidinyloxy group is in the 6-position of the quinazoline ring.
18. A quinazoline derivative according to any one of the preceding claims, wherein R³ is selected from hydrogen, (1-6C)alkyl, (3-6C)cycloalkyl, (3-6C)cycloalkyl(1-3C)alkyl (2-6C)alkanoyl;
30 and wherein any (1-6C)alkyl or (2-6C)alkanoyl group within R³ is optionally substituted by 1 or 2 substituents independently selected from halogeno, hydroxy and (1-6C)alkyl and/or optionally a substituent selected from cyano, nitro, (2-8C)alkenyl,

(2-8C)alkynyl, (1-6C)alkoxy and NR^cR^d , wherein R^c is hydrogen or (1-4C)alkyl and R^d is hydrogen or (1-4C)alkyl.

19. A quinazoline derivative according to anyone of the preceding claims, wherein R^3 is
5 methyl.

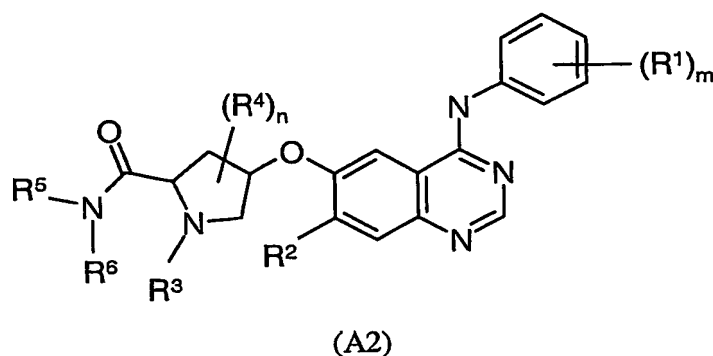
20. A quinazoline derivative according to anyone of the preceding claims, wherein n is 0, 1 or 2 and R^4 is independently selected from methyl, ethyl, methoxy, ethoxy, hydroxyl and oxo.

10 21. A quinazoline derivative according to anyone of the preceding claims, wherein n is 0.

22. A quinazoline derivative according to anyone of the preceding claims, wherein the $-\text{CONR}^5\text{R}^6$ group is in the 2-position of the pyrrolidine ring.

15 23. A quinazoline derivative according to anyone of the preceding claims, wherein the substituted-quinazolinyl group is in the 3-position of the pyrrolidine ring.

24. A quinazoline derivative according to any one of the preceding claims having a structural sub-formula A2



wherein:

m is 2 and R^1 is 2-fluoro and 3-chloro;

R^2 is methoxy;

25 R^3 is methyl;

n is 0;

and R⁵ is hydrogen or (1-6C)alkyl and R⁶ is selected from substituted-(1-6C)alkyl (wherein substituted-(1-6C)alkyl is (1-6C)alkyl substituted by 1, 2 or 3 substituents independently selected from (1-6C)alkoxycarbonyl, carbamoyl, (2-6C)alkanoylamino, and oxo or a (1-6C)alkoxycarbonyl together with a hydroxy group), (1-6C)alkoxy, (1-6C)alkylsulfonyl, (3-7)heterocyclyl (wherein the heterocyclyl is carbon linked), heteroaryl, (3-7)heterocyclyl(1-6C)alkyl (wherein the heterocyclyl is carbon linked to the (1-6C)alkyl moiety) and heteroaryl(1-6C)alkyl, and wherein any heteroaryl or (3-7)heterocyclyl group within R⁶ is optionally substituted (on any available carbon atoms) by 1, 2 or 3 substituents independently selected from halogeno, (1-6C)alkyl, hydroxy(1-6C)alkyl, (1-6C)alkoxycarbonyl, carbamoyl, (2-6C)alkanoylamino and hydroxy and/or optionally a substituent selected from oxo, cyano, nitro and (1-4C)alkoxy, and wherein any heteroaryl or heterocyclyl group within R⁶ is optionally substituted on any available ring nitrogen (provided the ring is not thereby quaternised) by (1-4C)alkyl or (2-4C)alkanoyl, or

15 R⁵ and R⁶ together with the nitrogen atom to which they are attached form a 4, 5 or 6 membered ring which contains one or two nitrogen atoms as the only heteroatoms present in the ring and which is optionally and which is substituted on an available ring carbon atom by 1 or 2 substituents independently selected from carbamoyl and (1-3C)alkylenedioxy.

20 25. A quinazoline derivative according to claim 24, wherein R⁶ is selected from (3-7)heterocyclyl (wherein the heterocyclyl is carbon linked), heteroaryl, (3-7)heterocyclyl(1-6C)alkyl (wherein the heterocyclyl is carbon linked to the (1-6C)alkyl moiety) and heteroaryl(1-6C)alkyl, and wherein any heteroaryl or (3-7)heterocyclyl group within R⁶ is optionally substituted (on any available carbon atoms) by 1, 2 or 3 substituents independently selected from halogeno, (1-6C)alkyl, hydroxy(1-6C)alkyl, (1-6C)alkoxycarbonyl, carbamoyl, (2-6C)alkanoylamino and hydroxy and/or optionally a substituent selected from oxo, cyano, nitro and (1-4C)alkoxy, and wherein any heteroaryl or heterocyclyl group within R⁶ is optionally substituted on any available ring nitrogen (provided the ring is not thereby quaternised) by (1-4C)alkyl or (2-4C)alkanoyl.

30

26. A quinazoline derivative selected from one or more of the following:

- (4*S*)-4-({4-[(3-chloro-2-fluorophenyl)amino]quinazolin-7-yl}oxy)-*N,N*,1-trimethyl-L-prolinamide;
- (4*S*)-4-({4-[(3-chloro-2-fluorophenyl)amino]quinazolin-7-yl}oxy)-1-methyl-L-prolinamide;
- 5 (4*S*)-4-({4-[(4-cyano-2-fluorophenyl)amino]-7-methoxyquinazolin-6-yl}oxy)-*N,N*,1-trimethyl-D-prolinamide;
- (4*S*)-4-({4-[(3-chloro-4-cyanophenyl)amino]-7-methoxyquinazolin-6-yl}oxy)-*N,N*,1-trimethyl-D-prolinamide;
- (4*S*)-4-[(4-{[3-chloro-4-(trifluoromethyl)phenyl]amino}-7-methoxyquinazolin-6-yl)oxy]-
- 10 *N,N*,1-trimethyl-D-prolinamide;
- (4*S*)-4-({4-[(5-chloropyridin-3-yl)amino]-7-methoxyquinazolin-6-yl}oxy)-*N,N*,1-trimethyl-D-prolinamide;
- (4*S*)-4-({4-[(2-fluoro-4-methylphenyl)amino]-7-methoxyquinazolin-6-yl}oxy)-*N,N*,1-trimethyl-D-prolinamide;
- 15 (4*S*)-4-({4-[(3-chloro-4-fluorophenyl)amino]-7-methoxyquinazolin-6-yl}oxy)-*N,N*,1-trimethyl-D-prolinamide;
- (4*S*)-4-({4-[(2-fluoro-4-hydroxyphenyl)amino]-7-methoxyquinazolin-6-yl}oxy)-*N,N*,1-trimethyl-D-prolinamide;
- (4*S*)-4-({4-[(2,4-difluorophenyl)amino]-7-methoxyquinazolin-6-yl}oxy)-*N,N*,1-trimethyl-D-
- 20 prolinamide;
- (4*S*)-4-({4-[(2,5-difluorophenyl)amino]-7-methoxyquinazolin-6-yl}oxy)-*N,N*,1-trimethyl-D-prolinamide;
- (4*S*)-4-({4-[(5-chloro-2-fluorophenyl)amino]-7-methoxyquinazolin-6-yl}oxy)-*N,N*,1-trimethyl-D-prolinamide;
- 25 (4*S*)-4-({4-[(4-chloro-2-fluorophenyl)amino]-7-methoxyquinazolin-6-yl}oxy)-*N,N*,1-trimethyl-D-prolinamide;
- (4*S*)-4-({4-[(5-chloro-2-hydroxyphenyl)amino]-7-methoxyquinazolin-6-yl}oxy)-*N,N*,1-trimethyl-D-prolinamide;
- (4*S*)-4-({4-[(3-chloro-4-methoxyphenyl)amino]-7-methoxyquinazolin-6-yl}oxy)-*N,N*,1-
- 30 trimethyl-D-prolinamide;
- (4*S*)-4-[(4-{[2-(aminosulfonyl)-5-chlorophenyl]amino}-7-methoxyquinazolin-6-yl)oxy]-*N,N*,1-trimethyl-D-prolinamide;

- (4S)-4-({7-methoxy-4-[(2,3,4-trifluorophenyl)amino]quinazolin-6-yl}oxy)-N,N,1-trimethyl-D-prolinamide;
- (4S)-4-[(4-{[2-fluoro-5-(trifluoromethyl)phenyl]amino}-7-methoxyquinazolin-6-yl)oxy]-N,N,1-trimethyl-D-prolinamide;
- 5 (4S)-4-[(4-{[2-fluoro-3-(trifluoromethyl)phenyl]amino}-7-methoxyquinazolin-6-yl)oxy]-N,N,1-trimethyl-D-prolinamide;
- (4S)-4-({4-[(3-chloro-2-methoxyphenyl)amino]-7-methoxyquinazolin-6-yl}oxy)-N,N,1-trimethyl-D-prolinamide;
- (4S)-4-({4-[(3-chloro-2-methylphenyl)amino]-7-methoxyquinazolin-6-yl}oxy)-N,N,1-trimethyl-D-prolinamide;
- 10 (4S)-4-({4-[(3-chloro-4-hydroxyphenyl)amino]-7-methoxyquinazolin-6-yl}oxy)-N,N,1-trimethyl-D-prolinamide;
- (4S)-4-({4-[(3-ethynylphenyl)amino]-7-methoxyquinazolin-6-yl}oxy)-N,N,1-trimethyl-D-prolinamide;
- 15 (4S)-4-({4-[(3-cyanophenyl)amino]-7-methoxyquinazolin-6-yl}oxy)-N,N,1-trimethyl-D-prolinamide;
- (4S)-4-({4-[(1*H*-indol-5-yl)amino]-7-methoxyquinazolin-6-yl}oxy)-N,N,1-trimethyl-D-prolinamide;
- (4S)-4-({4-[(3-chloro-1*H*-indol-5-yl)amino]-7-methoxyquinazolin-6-yl}oxy)-N,N,1-trimethyl-D-prolinamide;
- 20 (4S)-4-({4-[(3-chloro-2-fluorophenyl)amino]-7-methoxyquinazolin-6-yl}oxy)-N-cyclopropyl-1-methyl-D-prolinamide;
- (4S)-4-({4-[(3-chloro-2-fluorophenyl)amino]-7-methoxyquinazolin-6-yl}oxy)-N-(cyclopropylmethyl)-1-methyl-D-prolinamide;
- 25 (4S)-4-({4-[(3-chloro-2-fluorophenyl)amino]-7-methoxyquinazolin-6-yl}oxy)-N-(2-methoxyethyl)-1-methyl-D-prolinamide;
- (4S)-4-({4-[(3-chloro-2-fluorophenyl)amino]-7-methoxyquinazolin-6-yl}oxy)-N-cyclopentyl-1-methyl-D-prolinamide;
- (4S)-4-({4-[(3-chloro-2-fluorophenyl)amino]-7-methoxyquinazolin-6-yl}oxy)-N-cyclopentylmethyl-1-methyl-D-prolinamide;
- 30 (4S)-4-({4-[(3-chloro-2-fluorophenyl)amino]-7-methoxyquinazolin-6-yl}oxy)-N-(2-methoxyethyl)-N,1-dimethyl-D-prolinamide;

- (4*S*)-4-({4-[(3-chloro-2-fluorophenyl)amino]-7-methoxyquinazolin-6-yl}oxy)-*N*-methoxy-1-methyl-D-prolinamide;
- (4*S*)-4-({4-[(3-chloro-2-fluorophenyl)amino]-7-methoxyquinazolin-6-yl}oxy)-*N*-cyclohexyl-1-methyl-D-prolinamide;
- 5 (4*S*)-4-({4-[(3-chloro-2-fluorophenyl)amino]-7-methoxyquinazolin-6-yl}oxy)-1-methyl-*N*-(tetrahydro-2*H*-pyran-4-yl)-D-prolinamide; and
- (4*S*)-4-({4-[(3-chloro-4-fluorophenyl)amino]-6-methoxyquinazolin-7-yl}oxy)-*N,N*,1-trimethyl-L-prolinamide;
- (4*S*)-4-({4-[(3-chloro-2-fluorophenyl)amino]-7-methoxyquinazolin-6-yl}oxy)-*N*-[(1*R*)-1-
- 10 (hydroxymethyl)-3-methylbutyl]-1-methyl-D-prolinamide;
- (4*S*)-4-({4-[(3-chloro-2-fluorophenyl)amino]-7-methoxyquinazolin-6-yl}oxy)-*N*-[(1*S*)-1-(hydroxymethyl)-3-methylbutyl]-1-methyl-D-prolinamide;
- (4*S*)-4-({4-[(3-chloro-2-fluorophenyl)amino]-7-methoxyquinazolin-6-yl}oxy)-*N*-(3-furylmethyl)-1-methyl-D-prolinamide;
- 15 (4*S*)-4-({4-[(3-chloro-2-fluorophenyl)amino]-7-methoxyquinazolin-6-yl}oxy)-*N*-(2-furylmethyl)-1-methyl-D-prolinamide;
- (4*S*)-4-({4-[(3-chloro-2-fluorophenyl)amino]-7-methoxyquinazolin-6-yl}oxy)-1-methyl-*N*-[(5-methylisoxazol-3-yl)methyl]-D-prolinamide;
- (4*S*)-4-({4-[(3-chloro-2-fluorophenyl)amino]-7-methoxyquinazolin-6-yl}oxy)-*N*-[2-(1*H*-
- 20 imidazol-1-yl)ethyl]-1-methyl-D-prolinamide;
- (2*S*)-1-[(4*S*)-4-({4-[(3-chloro-2-fluorophenyl)amino]-7-methoxyquinazolin-6-yl}oxy)-1-methyl-D-prolyl]azetidine-2-carboxamide;
- (4*S*)-4-({4-[(3-chloro-2-fluorophenyl)amino]-7-methoxyquinazolin-6-yl}oxy)-*N*-[(2*R*)-2,3-dihydroxypropyl]-1-methyl-D-prolinamide;
- 25 (4*S*)-4-({4-[(3-chloro-2-fluorophenyl)amino]-7-methoxyquinazolin-6-yl}oxy)-1-methyl-*N*-(1-methyl-1*H*-pyrazol-5-yl)-D-prolinamide;
- (4*S*)-4-({4-[(3-chloro-2-fluorophenyl)amino]-7-methoxyquinazolin-6-yl}oxy)-1-methyl-*N*-3-thienyl-D-prolinamide; and
- (4*S*)-4-({4-[(3-chloro-2-fluorophenyl)amino]-7-methoxyquinazolin-6-yl}oxy)-1-methyl-*N*-(3-
- 30 methyl-1*H*-pyrazol-5-yl)-D-prolinamide;
- methyl (4*S*)-4-({4-[(3-chloro-2-fluorophenyl)amino]-7-methoxyquinazolin-6-yl}oxy)-1-methyl-D-prolyl-L-serinate;

- (4*S*)-4-({4-[(3-chloro-2-fluorophenyl)amino]-7-methoxyquinazolin-6-yl}oxy)-*N*-(2-hydroxy-1,1-dimethylethyl)-1-methyl-D-prolinamide;
- (4*S*)-4-({4-[(3-chloro-2-fluorophenyl)amino]-7-methoxyquinazolin-6-yl}oxy)-1-methyl-D-prolylglycinamide;
- 5 (4*S*)-*N*-[2-(acetylamino)ethyl]-4-({4-[(3-chloro-2-fluorophenyl)amino]-7-methoxyquinazolin-6-yl}oxy)-1-methyl-D-prolinamide;
- (4*S*)-4-({4-[(3-chloro-2-fluorophenyl)amino]-7-methoxyquinazolin-6-yl}oxy)-*N*-[(3*S*,4*R*)-4-hydroxytetrahydrofuran-3-yl]-1-methyl-D-prolinamide;
- (4*S*)-4-({4-[(3-chloro-2-fluorophenyl)amino]-7-methoxyquinazolin-6-yl}oxy)-*N*-[1-
- 10 (hydroxymethyl)cyclopentyl]-1-methyl-D-prolinamide;
- (4*S*)-4-({4-[(3-chloro-2-fluorophenyl)amino]-7-methoxyquinazolin-6-yl}oxy)-*N*-[(1*S*)-1-(hydroxymethyl)-2-methylpropyl]-1-methyl-D-prolinamide;
- (4*S*)-4-({4-[(3-chloro-2-fluorophenyl)amino]-7-methoxyquinazolin-6-yl}oxy)-*N*-[2-(1*H*-imidazol-4-yl)ethyl]-1-methyl-D-prolinamide;
- 15 (4*S*)-4-({4-[(3-chloro-2-fluorophenyl)amino]-7-methoxyquinazolin-6-yl}oxy)-*N*-(2-methoxy-1-methylethyl)-1-methyl-D-prolinamide;
- (4*S*)-4-({4-[(3-chloro-2-fluorophenyl)amino]-7-methoxyquinazolin-6-yl}oxy)-1-methyl-*N*-(2,2,2-trifluoroethyl)-D-prolinamide;
- (4*S*)-*N*-allyl-4-({4-[(3-chloro-2-fluorophenyl)amino]-7-methoxyquinazolin-6-yl}oxy)-1-
- 20 methyl-D-prolinamide;
- (4*S*)-4-({4-[(3-chloro-2-fluorophenyl)amino]-7-methoxyquinazolin-6-yl}oxy)-*N*-(2-ethoxyethyl)-1-methyl-D-prolinamide;
- (4*S*)-4-({4-[(3-chloro-2-fluorophenyl)amino]-7-methoxyquinazolin-6-yl}oxy)-*N*-(4-hydroxycyclohexyl)-1-methyl-D-prolinamide;
- 25 (4*S*)-4-({4-[(3-chloro-2-fluorophenyl)amino]-7-methoxyquinazolin-6-yl}oxy)-1-methyl-*N*-(2-methylprop-2-en-1-yl)-D-prolinamide;
- (4*S*)-4-({4-[(3-chloro-2-fluorophenyl)amino]-7-methoxyquinazolin-6-yl}oxy)-*N*-[(1*S*)-1-(hydroxymethyl)propyl]-1-methyl-D-prolinamide;
- (4*S*)-4-({4-[(3-chloro-2-fluorophenyl)amino]-7-methoxyquinazolin-6-yl}oxy)-*N*-[(2*S*)-2,3-
- 30 dihydroxypropyl]-1-methyl-D-prolinamide;
- (4*S*)-4-({4-[(3-chloro-2-fluorophenyl)amino]-7-methoxyquinazolin-6-yl}oxy)-*N*-(1*H*-imidazol-2-ylmethyl)-1-methyl-D-prolinamide;

- (4*S*)-4-({4-[(3-chloro-2-fluorophenyl)amino]-7-methoxyquinazolin-6-yl}oxy)-*N*-[2-(2-furyl)ethyl]-1-methyl-D-prolinamide;
- (4*S*)-4-({4-[(3-chloro-2-fluorophenyl)amino]-7-methoxyquinazolin-6-yl}oxy)-1-methyl-*N*-(tetrahydro-2*H*-pyran-4-ylmethyl)-D-prolinamide;
- 5 (4*S*)-4-({4-[(3-chloro-2-fluorophenyl)amino]-7-methoxyquinazolin-6-yl}oxy)-*N*-[(1*S*)-2-hydroxy-1-methylethyl]-1-methyl-D-prolinamide;
- (4*S*)-4-({4-[(3-chloro-2-fluorophenyl)amino]-7-methoxyquinazolin-6-yl}oxy)-*N*-[(1*R*)-2-hydroxy-1-methylethyl]-1-methyl-D-prolinamide;
- (4*S*)-4-({4-[(3-chloro-2-fluorophenyl)amino]-7-methoxyquinazolin-6-yl}oxy)-*N*-[(2*R*)-2-
- 10 hydroxypropyl]-1-methyl-D-prolinamide;
- (4*S*)-4-({4-[(3-chloro-2-fluorophenyl)amino]-7-methoxyquinazolin-6-yl}oxy)-*N*-[(2*S*)-2-hydroxypropyl]-1-methyl-D-prolinamide;
- (4*S*)-4-({4-[(3-chloro-2-fluorophenyl)amino]-7-methoxyquinazolin-6-yl}oxy)-1-methyl-*N*-[(2*R*)-tetrahydrofuran-2-ylmethyl]-D-prolinamide;
- 15 (4*S*)-4-({4-[(3-chloro-2-fluorophenyl)amino]-7-methoxyquinazolin-6-yl}oxy)-1-methyl-*N*-[(2*S*)-tetrahydrofuran-2-ylmethyl]-D-prolinamide
- N*-(3-chloro-2-fluorophenyl)-7-methoxy-6-{{(3*S*,5*R*)-1-methyl-5-(pyrrolidin-1-ylcarbonyl)pyrrolidin-3-yl}oxy}quinazolin-4-amine;
- N*-(3-chloro-2-fluorophenyl)-7-methoxy-6-{{(3*S*,5*R*)-1-methyl-5-[(4-methylpiperazin-1-
- 20 yl)carbonyl]pyrrolidin-3-yl}oxy}quinazolin-4-amine
- 6-{{(3*S*,5*R*)-5-(azetidin-1-ylcarbonyl)-1-methylpyrrolidin-3-yl}oxy}-*N*-(3-chloro-2-fluorophenyl)-7-methoxyquinazolin-4-amine;
- (4*S*)-4-({4-[(3-chloro-2-fluorophenyl)amino]-7-methoxyquinazolin-6-yl}oxy)-*N*-(cyanomethyl)-*N*,1-dimethyl-D-prolinamide;
- 25 (4*S*)-4-({4-[(3-chloro-2-fluorophenyl)amino]-7-methoxyquinazolin-6-yl}oxy)-*N*-(cyanomethyl)-1-methyl-D-prolinamide;
- (4*S*)-4-({4-[(3-chloro-2-fluorophenyl)amino]-7-methoxyquinazolin-6-yl}oxy)-*N*,1-dimethyl-*N*-[(2*S*)-2-pyrrolidin-1-ylpropyl]-D-prolinamide;
- (4*S*)-4-({4-[(3-chloro-2-fluorophenyl)amino]-7-methoxyquinazolin-6-yl}oxy)-*N*-[(1*R*)-2-
- 30 hydroxy-1-methylethyl]-*N*,1-dimethyl-D-prolinamide;
- (4*S*)-4-({4-[(3-chloro-2-fluorophenyl)amino]-7-methoxyquinazolin-6-yl}oxy)-*N*,1-dimethyl-*N*-(1-methylpiperidin-4-yl)-D-prolinamide;

- (4*S*)-4-({4-[(3-chloro-2-fluorophenyl)amino]-7-methoxyquinazolin-6-yl}oxy)-*N*,1-dimethyl-*N*-(tetrahydro-2*H*-pyran-4-yl)-*D*-prolinamide;
- (4*R*)-4-({4-[(3-chloro-2-fluorophenyl)amino]-7-methoxyquinazolin-6-yl}oxy)-1-methyl-*N*-prop-2-yn-1-yl-*L*-prolinamide;
- 5 1-[[[(2*S*,4*R*)-4-[[4-[(3-chloro-2-fluorophenyl)amino]-7-methoxy-6-quinazolinyl]oxy]-1-methyl-2-pyrrolidinyl]carbonyl]-3-pyrroline;
- (4*R*)-4-({4-[(3-chloro-2-fluorophenyl)amino]-7-methoxyquinazolin-6-yl}oxy)-*N*-(cyanomethyl)-1-methyl-*L*-prolinamide;
- (4*R*)-4-({4-[(3-chloro-2-fluorophenyl)amino]-7-methoxyquinazolin-6-yl}oxy)-*N*-(2-
- 10 cyanoethyl)-1-methyl-*L*-prolinamide;
- (4*R*)-4-({4-[(3-chloro-2-fluorophenyl)amino]-7-methoxyquinazolin-6-yl}oxy)-*N*-(cyanomethyl)-*N*,1-dimethyl-*L*-prolinamide;
- (4*R*)-4-({4-[(3-chloro-2-fluorophenyl)amino]-7-methoxyquinazolin-6-yl}oxy)-*N*-(2-methoxyethyl)-1-methyl-*L*-prolinamide;
- 15 (4*R*)-4-({4-[(3-chloro-2-fluorophenyl)amino]-7-methoxyquinazolin-6-yl}oxy)-*N*-cyclopropyl-1-methyl-*L*-prolinamide;
- (4*R*)-4-({4-[(3-chloro-2-fluorophenyl)amino]-7-methoxyquinazolin-6-yl}oxy)-*N*-cyclopentyl-1-methyl-*L*-prolinamide;
- N*-(3-chloro-2-fluorophenyl)-7-methoxy-6-({[(3*R*,5*S*)-1-methyl-5-[(4-methylpiperazin-1-
- 20 yl)carbonyl]pyrrolidin-3-yl}oxy)quinazolin-4-amine;
- (3*S*)-1-[(4*R*)-4-({4-[(3-chloro-2-fluorophenyl)amino]-7-methoxyquinazolin-6-yl}oxy)-1-methyl-*L*-prolyl]pyrrolidin-3-ol
- (4*R*)-4-({4-[(3-chloro-2-fluorophenyl)amino]-7-methoxyquinazolin-6-yl}oxy)-*N*-(cyclopropylmethyl)-1-methyl-*L*-prolinamide;
- 25 (4*R*)-4-({4-[(3-chloro-2-fluorophenyl)amino]-7-methoxyquinazolin-6-yl}oxy)-*N*-cyclohexyl-*N*,1-dimethyl-*L*-prolinamide;
- (4*R*)-4-({4-[(3-chloro-2-fluorophenyl)amino]-7-methoxyquinazolin-6-yl}oxy)-1-methyl-*N*-(tetrahydro-2*H*-pyran-4-yl)-*L*-prolinamide;
- N*-(3-chloro-2-fluorophenyl)-7-methoxy-6-{{[(3*R*,5*S*)-1-methyl-5-(pyrrolidin-1-
- 30 ylcarbonyl]pyrrolidin-3-yl}oxy}quinazolin-4-amine;
- (4*R*)-4-({4-[(3-chloro-2-fluorophenyl)amino]-7-methoxyquinazolin-6-yl}oxy)-*N*-(2-hydroxyethyl)-*N*,1-dimethyl-*L*-prolinamide;

- (4R)-4-({4-[(3-chloro-2-fluorophenyl)amino]-7-methoxyquinazolin-6-yl}oxy)-N-[2-(dimethylamino)ethyl]-1-methyl-L-prolinamide;
- (4R)-4-({4-[(3-chloro-2-fluorophenyl)amino]-7-methoxyquinazolin-6-yl}oxy)-N,1-dimethyl-N-(1-methylpiperidin-4-yl)-L-prolinamide;
- 5 6-({(3R,5S)-5-[(4-acetylpiperazin-1-yl)carbonyl]-1-methylpyrrolidin-3-yl}oxy)-N-(3-chloro-2-fluorophenyl)-7-methoxyquinazolin-4-amine;
- 1-[(4R)-4-({4-[(3-chloro-2-fluorophenyl)amino]-7-methoxyquinazolin-6-yl}oxy)-1-methyl-L-prolyl]piperidin-4-ol;
- (4R)-4-({4-[(3-chloro-2-fluorophenyl)amino]-7-methoxyquinazolin-6-yl}oxy)-N-(2-methoxyethyl)-N,1-dimethyl-L-prolinamide;
- 10 (4R)-4-({4-[(3-chloro-2-fluorophenyl)amino]-7-methoxyquinazolin-6-yl}oxy)-N-cyclohexyl-1-methyl-L-prolinamide;
- (4S)-4-({4-[(3-chloro-2-fluorophenyl)amino]-7-methoxyquinazolin-6-yl}oxy)-N-cyclopropyl-1-methyl-L-prolinamide;
- 15 (4S)-4-({4-[(3-chloro-2-fluorophenyl)amino]-7-methoxyquinazolin-6-yl}oxy)-N-(2-methoxyethyl)-1-methyl-L-prolinamide;
- (4S)-4-({4-[(3-chloro-2-fluorophenyl)amino]-7-methoxyquinazolin-6-yl}oxy)-N-cyclohexyl-N,1-dimethyl-L-prolinamide;
- (4S)-4-({4-[(3-chloro-2-fluorophenyl)amino]-7-methoxyquinazolin-6-yl}oxy)-1-methyl-N-
- 20 (tetrahydro-2H-pyran-4-yl)-L-prolinamide;
- (4S)-4-({4-[(3-chloro-2-fluorophenyl)amino]-7-methoxyquinazolin-6-yl}oxy)-N-(2-methoxyethyl)-N,1-dimethyl-L-prolinamide;
- (4S)-4-({4-[(3-chloro-2-fluorophenyl)amino]-7-methoxyquinazolin-6-yl}oxy)-N,1-dimethyl-N-(1-methylpiperidin-4-yl)-L-prolinamide;
- 25 (4S)-4-({4-[(3-chloro-2-fluorophenyl)amino]-7-methoxyquinazolin-6-yl}oxy)-N-cyclopentyl-1-methyl-L-prolinamide;
- (4S)-4-({4-[(3-chloro-2-fluorophenyl)amino]-7-methoxyquinazolin-6-yl}oxy)-N-methoxy-1-methyl-L-prolinamide;
- (4S)-4-({4-[(3-chloro-2-fluorophenyl)amino]-7-methoxyquinazolin-6-yl}oxy)-N-
- 30 (cyclopropylmethyl)-1-methyl-L-prolinamide;
- (4S)-4-({4-[(3-chloro-2-fluorophenyl)amino]-7-methoxyquinazolin-6-yl}oxy)-N-cyclohexyl-1-methyl-L-prolinamide;

and pharmaceutically-acceptable salts thereof.

27. (4S)-4-({4-[(3-chloro-2-fluorophenyl)amino]quinazolin-7-yl}oxy)-1-methyl-L-prolinamide trifluoroacetic acid salt.

5

28. A pharmaceutical composition which comprises a quinazoline derivative of the Formula I, or a pharmaceutically-acceptable salt or prodrug form thereof, as defined in any one of claims 1 to 25 in association with a pharmaceutically-acceptable diluent or carrier.

10 29. A quinazoline derivative of the Formula I as defined in any one of claims 1 to 25, or a pharmaceutically acceptable salt or prodrug form thereof, for use as a medicament.

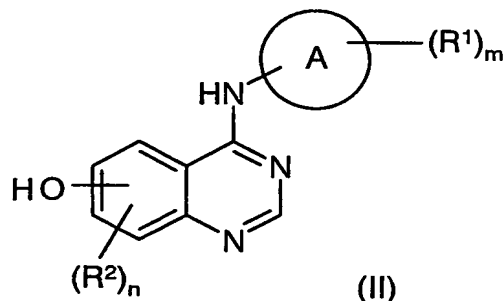
30. The use of a quinazoline derivative of the Formula I, or a pharmaceutically-acceptable salt or prodrug form thereof, as defined in any one of claims 1 to 25 in the manufacture of a
15 medicament for use in the production of an anti-proliferative effect in a warm-blooded animal.

31. A method for producing an anti-proliferative effect in a warm-blooded animal in need of such treatment, which comprises administering to, said animal a quinazoline derivative of the Formula I, or a pharmaceutically acceptable salt or prodrug form thereof, as defined in any
20 one of claims 1 to 25.

32. A process for the preparation of a quinazoline derivative of the Formula I as defined in Claim 1 which is selected from one of the following:

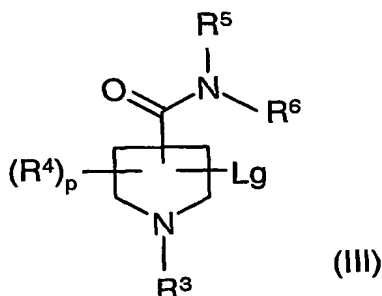
Process (a) reacting a compound of the Formula II:

25



wherein R^1 , R^2 , A, m and n have any of the meanings defined in claim 1, except that any functional group is protected if necessary,

with a compound of the Formula III in the presence of a suitable base:



5

wherein R^3 , R^4 , R^5 , R^6 and p have any of the meanings defined in claim 1, except that any functional group is protected if necessary and Lg is a displaceable group,

and whereafter any protecting group that is present is removed;

- 10 **Process (b)** modifying a substituent in, or introducing a substituent into, another quinazoline derivative of Formula I, or a pharmaceutically acceptable salt thereof, as defined in claim 1, except that any functional group is protected if necessary,

and whereafter any protecting group that is present is removed;

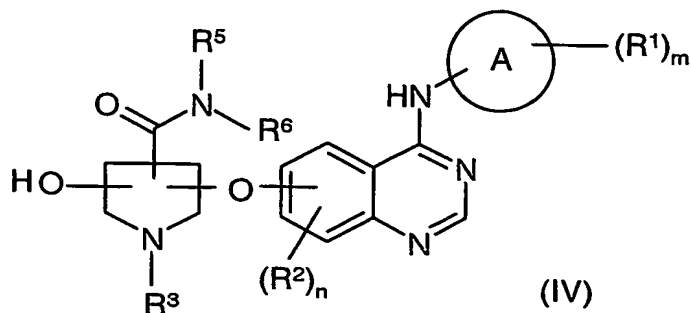
- Process (c)** the removal of a protecting group from a quinazoline derivative of Formula I,
15 or a pharmaceutically acceptable salt thereof, as claimed in claim 1;

Process (d) reacting a compound of the Formula II as defined in reference to process (a) above with a compound of the Formula III as defined in reference to process (a) above, except Lg is OH, under Mitsunobu conditions, and whereafter any protecting group that is present is removed by conventional means;

- 20 **Process (e)** For the preparation of those compounds of the Formula I defined in claim 1 wherein R^4 is a hydroxy group, by the cleavage of a quinazoline derivative of the Formula I wherein R^4 is a (1-4C)alkoxy group.

Process (f) For the preparation of those compounds of the Formula I defined in claim 1 wherein R^4 is (1-4C)alkoxy, by the reaction of a compound of the Formula IV:

25



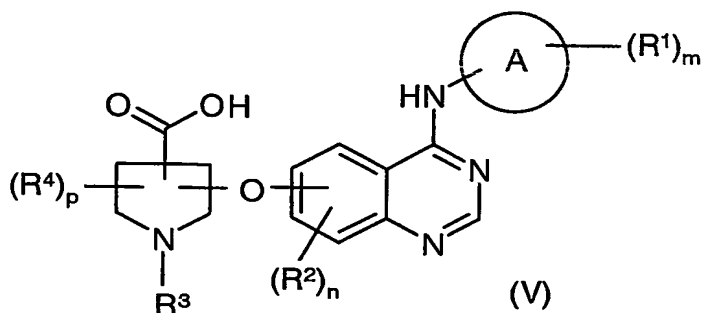
with a compound of the formula (1-4C)alkyl-Lg in the presence of a base, wherein Lg is a displaceable group,

5 and whereafter any protecting group that is present is removed by conventional means;

Process (g) For the preparation of those compounds of the Formula I defined in claim 1 wherein R¹, R², R⁴ or R⁶ contain a (1-6C)alkoxy or substituted (1-6C)alkoxy group or a (1-6C)alkylamino or substituted (1-6C)alkylamino group, said process comprising the alkylation of a quinazoline derivative of the Formula I wherein R¹, R², R⁴ or R⁶ contain a hydroxy group

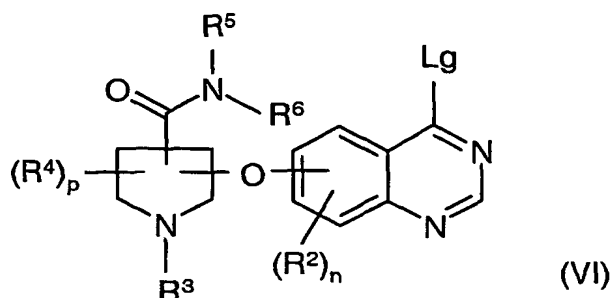
10 or a primary or secondary amino group as appropriate;

Process (h) reacting a compound of the formula (V) or reactive derivative thereof

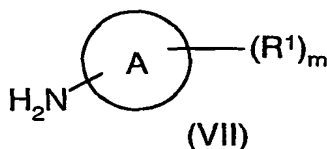


15 with a compound of the formula HNR⁵R⁶ or a suitable salt thereof in the presence of a base and in an inert solvent;

Process (i) reacting a compound of the formula VI:



wherein R^1 , R^2 , R^3 , R^4 , R^5 , R^6 , n and p , have any of the meanings defined in claim 1, except that any functional group is protected if necessary, and Lg is a displaceable group as defined in
 5 reference to Process (a) above,
 with an aniline of the formula VII in the presence of a suitable acid:



wherein R^1 and m have any of the meanings defined in claim 1, except that any
 10 functional group is protected if necessary,

Process (j) Forming the group $-\text{CON}(R^5)R^6$ by reacting to the corresponding carboxy compound, wherein any functional groups are protected if necessary, with a primary or secondary amine or a heterocyclic group containing an NH group;

and whereafter any protecting group that is present is removed by conventional means.